L Number	Hits	Search Text	T D D	
1	3076	knockout and promoter and operably	DB	Time stamp
	50,0	knockout and promoter and operably	USPAT;	2003/06/04 08:0
2	2000		US-PGPUB	00:0
۷	3020	( this street and promoter and operation and	USPAT;	2002/06/04 00 0
İ		(operably adj linked)		2003/06/04 08:0
3	2913	((knockout and promoter and operably) and	US-PGPUB	
		(charable and promoter and operably) and	USPAT;	2003/06/04 08:0
4	_	(operably adj linked)) and mouse	US-PGPUB	
1	ا م	(((knockout and promoter and operably) and	USPAT;	2003/06/04 08:0
		(operably adj linked)) and mouse) and	US-PGPUB	2003/00/04 08:0
		(promoter adj fusion)	US-FGFUB	

FILE 'MEDLINE, BIOSIS, JUN 2003	EMBASE,	LIFESCI,	CAPLUS'	ENTERED	АТ	06:39:57	ON	04
2 S RAMP1 AND	KNOCKOU'	T						
2 DUP REM L1	(O DUPLIC	CATES REMO	びだり					
60 S RAMP1 AND	TRANSMEN	OFFED TOTAL	J V E D J					
22 DUP REM L3	(38 DUPL	TCATES REA	MOVED)					
1 S RAMP1 AND	DEFICIEN	TOTTLE KEL	TOVED)					
231 S RAMP1 AND								
19 S L6 AND RE		r						
13 DUP REM L7			יעבים ו					
	(0 501111	STILLS KEINC	) V E D )					

L1 L2 L3 L4 L5

L6 L7

 $^{\text{L8}}$ 

L Number	Hits	Search Text	DB	Time stamp
2	6	RAMP1 and transmembrane	USPAT; US-PGPUB;	2003/06/04 06:53
1	31	RAMP1	DERWENT USPAT; US-PGPUB; DERWENT	2003/06/04 07:07

ANSWER 1 OF 13 CAPLUS COPYRIGHT 2003 ACS 8 AN 2003:261950 CAPLUS DN 138:282345 Humanized calcitonin gene-related peptide (CGRP) receptor comprising TIcalcitonin-receptor-like receptor (CRLR) and the receptor-activitymodifying protein 1 (RAMP1) IN Kane, Stefanie A.; Salvatore, Christopher A.; Mallee, John J.; Koblan, Kenneth S.; Oliver, Kevin R. PΑ Merck & Co., Inc., USA PCT Int. Appl., 60 pp. SO CODEN: PIXXD2 DTPatent LA English FAN.CNT 1 KIND DATE APPLICATION NO. DATE PATENT NO. \_\_\_\_\_ PΙ WO 2003027252 A2 20030403 WO 2002-US30501 20020926 W: CA, JP, US RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR PRAI US 2001-325295P Ρ 20010927 The invention provides method for creating a humanized version of a calcitonin gene-related peptide (CGRP) receptor, which comprises the G-protein coupled receptor calcitonin-receptor-like receptor (CRLR) and the receptor-activity-modifying protein 1 (RAMP1). The humanized CGRP receptors of the present invention attain pharmacol. profiles similar to the wild type human receptor via modifications to the resp. mammalian RAMP1 nucleotide sequence, specifically at amino acid 74. Also described are related recombinant vectors, recombinant hosts and assocd. methods for generating such humanized CGRP receptors. Also presented are non-human transgenic animals which express humanized RAMP1. Such animals have been

engineered to provide for a CGRP pharmacol. profile similar to human CGRP. Antagonist of CGRP function may be useful in the treatment of various disorders such as migraine headaches, pain indications, menopausal hot flashes, migraine prophylaxis, chronic tension type headache, cluster headache, neurogenic or chronic inflammation, gastrointestinal disorders,

type 2 diabetes and cardiovascular disorders.